09/368,989 L/Cook 3/11/05-

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(FILE 'HOME' ENTERED AT 10:08:29 ON 11 MAR 2005)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, CANCERLIT, JAPIO' ENTERED AT 10:08:52 ON 11 MAR 2005

L1 3460 S (ANTIGEN BINDING SITE)

0 S (COMPLEMENTARY DETERMING SEG?)

L3 2 S (COMPLEMENTARY DETERMINING SEG?)

0 S L1 AND L2

L5 2376516 S ANTIBOD?

93 S L5 AND (180 DEGREE)

L7 13825 S L5 AND (HYDROPHOBIC?)

L8 3 S L6 AND L7

L9 3 DUPLICATE REMOVE L8 (0 DUPLICATES REMOVED)

=>

L2

L4

L6

ANSWER 1 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN

- AN 2003:184630 CAPLUS
- ED Entered STN: 11 Mar 2003
- TI A geomorphic classification of antibody binding sites
- AU Houk, K. N.; Lee, Michelle; Schallhorn, Julie; Sugimoto, Keiki; Leach, Andrew G.
- CS Department of Chemistry and Biochemistry, University of California, Los Angeles, CA, 90095-1569, USA
- SO Abstracts of Papers, 225th ACS National Meeting, New Orleans, LA, United States, March 23-27, 2003 (2003), ORGN-135 Publisher: American Chemical Society, Washington, D. C. CODEN: 69DSA4
- DT Conference; Meeting Abstract
- LA English
- AB The topogs. of antibody binding sites have been classified in analogy to familiar geomorphic features of our surroundings on earth. 265 available antibody crystal structures from the Protein Data Bank were analyzed according to this scheme. The binding site topog. classification is a modified scheme based on previous schemes but using readily recognizable geomorphic descriptors. There are five categories: cave and crater (mostly hapten binders), canyon and valley (mostly peptide and carbohydrate binders) and plain (mostly protein binders). Known catalytic antibody structures are mainly of the cave type, providing a deep hydrophobic pocket for binding of organic mol. substrates and transition states. The anal. was carried out at the UCLA Visualization Portal, where the 3D structures were analyzed visually by manipulation and assessment of the surface renderings of each crystal structure with a 3D projector in a 180 degrees concave viewing facility.

```
2003:660251 CAPLUS
AN
DN
     139:281226
ED
     Entered STN: 25 Aug 2003
     Preparation of core/shell type composite particles of hydroxyapatite and
ТT
     liposome
     Chu, Maoquan; Xu, Yuhong; Liu, Shupeng
IN
     Shanghai Jiaotong University, Peop. Rep. China
PΑ
     Faming Zhuanli Shenging Gongkai Shuomingshu, 7 pp.
SO
     CODEN: CNXXEV
DT
     Patent
     Chinese
LΑ
IC
     ICM A61K047-02
     ICS A61K047-46; B82B001-00; A61L027-12; A61P035-00
CC
     63-6 (Pharmaceuticals)
FAN.CNT 1
     PATENT NO.
                               DATE
                       KIND
                                          APPLICATION NO.
                       ----
    CN 1374132
                                                                  20020322
                        Α
                               20021016 CN 2002-111131
PRAI CN 2002-111131
                               20020322
CLASS
             CLASS PATENT FAMILY CLASSIFICATION CODES
PATENT NO.
 ______
               ICM A61K047-02
 CN 1374132
                       A61K047-46; B82B001-00; A61L027-12; A61P035-00
                ICS
     The core/shell type composite particle used as drug carrier is prepared by
AB
     preparing liposome membrane from phospholipid, lysolipid, glycolipid, sterol,
     cationic lipid, detergent, and/or amphoteric polymer via film coating
     method; encapsulating one or more of ions solution (pH 0-14) or
     hydroxyapatite-forming substance Ca3(PO4)2, CaO, Ca4P2O9, NaOH, and/or
     NH4OH at 0-100^{\circ}, and/or freezing at 0-(-180).
     degree. and melting at 0-100° 0-20 times; removing the un-
     encapsulated ions or hydroxyapatite-forming substance via column
     chromatog ; and allowing to react with another kinds of ions . The
     core/shell type composite particle may be modified with hydrophilic or
     hydrophobic polymer, ligand, antibody, cytokine,
     peptide, and/or nuclei acid during film-formation process or after
     formation of the core/shell type composite particle.
ST
     composite particle liposome hydroxyapatite drug carrier
ΙT
     Polymer morphology
        (core-shell; preparation of core/shell type composite particles of
       hydroxyapatite and liposome)
IT
     Drug delivery systems
        (liposomes; preparation of core/shell type composite particles of
       hydroxyapatite and liposome)
IT
    Lime (chemical)
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (preparation of core/shell type composite particles of hydroxyapatite and
        liposome)
IT
    1306-06-5, Hydroxyapatite 1310-58-3, Potassium hydroxide, biological
              1310-73-2, Sodium hydroxide, biological studies 7664-41-7,
    Ammonia, biological studies 7757-93-9, Calcium hydrogen phosphate 7758-87-4, Calcium phosphate 7790-76-3, Calcium pyrophosphate
    10043-52-4, Calcium chloride, biological studies 10124-37-5, Calcium
    nitrate
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (preparation of core/shell type composite particles of hydroxyapatite and
       liposome)
```

ANSWER 2 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN

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ANSWER 3 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN
     1997:385684 CAPLUS
AN
     127:9125
DN
     Entered STN: 21 Jun 1997
ED
ΤI
     Thermosensitive biodegradable polymers based on poly(ether-ester) block
     copolymers
     Cha, Younsik; Choi, Young Kweon; Bae, You Han
IN
     Macromed, Inc., USA
PA
SO
     PCT Int. Appl., 46 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
     ICM A61K009-10
IC
CC
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 35, 38
     PATENT NO.
                       KIND
                                DATE
                                          APPLICATION NO. DATE
                        ____
PΙ
     WO 9715287
                        A1
                               19970501 WO 1996-US17023 19961025
         W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GE, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT,
             RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, UZ, VN, AM,
            AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI
                                         US 1995-548185
     US 5702717
                         Α
                                19971230
                                                                   19951025
     ZA 9608944
                         Α
                                19970721
                                           ZA 1996-8944
                                                                   19961024
     CA 2235413
                         AA
                                19970501
                                           CA 1996-2235413
                                                                  19961025
     AU 9675200
                         A1
                                19970515
                                           AU 1996-75200
                                                                   19961025
     EP 863745
                         A1
                                19980916
                                           EP 1996-937727
                                                                   19961025
     EP 863745
                         B1
                                20040526
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
                         T2
     JP 11513985
                                19991130
                                           JP 1996-516762
                                                                   19961025
     AT 267584
                         Ε
                                20040615
                                           AT 1996-937727
                                                                   19961025
PRAI US 1995-548185
                        Α
                                19951025
     WO 1996-US17023
                         W
                                19961025
CLASS
                CLASS PATENT FAMILY CLASSIFICATION CODES
                ----
                       ICM
 WO 9715287
                       A61K009-10
 WO 9715287
                ECLA
                       A61K009/00M4; A61K038/18G; A61K038/20B; A61K038/22G;
                       A61K038/28; A61K047/34
 US 5702717
                ECLA
                       A61K009/00M4; A61K038/22G; A61K038/28; A61K047/34;
                       A61K038/18G; A61K038/20B
AB
    A system and method for the parenteral delivery of a drug in a
    biodegradable polymeric matrix to a warm blood animal as a liquid with the
     resultant formation of a gel depot for the controlled release of the drug.
     The system comprises an injectable biodegradable block copolymeric drug
     delivery liquid having reverse thermal gelation properties. The liquid is an
     aqueous solution having dissolved or dispersed therein an effective amount of a
     drug intimately contained in a biodegradable block copolymer matrix. The
     copolymer has a reverse gelation temperature below the body temperature of the
animal
     to which it is administered and is made of of (1) a hydrophobic
    A polymer block comprising a member selected from the group consisting of
    poly(\alpha-hydroxy\ acids) and poly(ethylene\ carbonates) and (2) a
```

A polymer block comprising a member selected from the group consisting of $poly(\alpha-hydroxy\ acids)$ and $poly(ethylene\ carbonates)$ and (2) a **hydrophobic** B polymer block comprising a polyethylene glycol. The liquid is stored below the reverse gelation temperature and is parenterally administered into the animal by i.m., i.p., s.c. or similar injection. Malic acid was polymerized with D,L-lactic acid to form a carboxyl group-containing oligomeric polyester. To melt of this carboxylated copolymer

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and then further heated at 180.degree. under a
     nitrogen atmospheric for 15 h to obtain the block copolymer of the invention.
     The above block copolymer wa dissolved in water and mixed with a solution of
     basic, heat stable platelet-derived growth factor. The drug was
     incorporated into the copolymer by a simultaneous dispersion and precipitation
     process. The precipitated copolymer containing drug particles was collected
and
     freeze dried.
     thermosensitive biodegradable polymer polyetherester block copolymer;
ST
     platelet derived growth factor parenteral pharmaceutical; malate lactide
     PEG block copolymer prepn
IT
     Antibodies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (monoclonal; thermosensitive biodegradable polymers based on
        poly(ether-ester) block copolymers)
     Drug delivery systems
ΙT
        (parenterals; thermosensitive biodegradable polymers based on
        poly(ether-ester) block copolymers)
IT
     Polyoxyalkylenes, biological studies
     Polyoxyalkylenes, biological studies
     RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); USES (Uses)
        (polyester-, block; thermosensitive biodegradable polymers based on
        poly(ether-ester) block copolymers)
     Polyesters, biological studies
IT
     Polyesters, biological studies
     RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); USES (Uses)
        (polyoxyalkylene-, block; thermosensitive biodegradable polymers based
        on poly(ether-ester) block copolymers)
IT
    Antitumor agents
     Enkephalins
     Interleukin 2
     Platelet-derived growth factors
     Tumor necrosis factors
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (thermosensitive biodegradable polymers based on poly(ether-ester)
        block copolymers)
IT
     Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\alpha; thermosensitive biodegradable polymers based on
        poly(ether-ester) block copolymers)
TΤ
     Interferons
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\beta); thermosensitive biodegradable polymers based on
        poly(ether-ester) block copolymers)
IT
     Interferons
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\gamma); thermosensitive biodegradable polymers based on
        poly(ether-ester) block copolymers)
ΙT
    190191-84-5P
    RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
    study); PREP (Preparation); USES (Uses)
        (multiblock; thermosensitive biodegradable polymers based on
        poly(ether-ester) block copolymers)
TТ
    113497-67-9P 115358-21-9P
                                  149479-29-8P
                                                  190191-83-4P
                                                                  190191-85-6DP,
    diol derivs.
                    190191-86-7P
                                   190191-87-8P
                                                  190191-88-9P
                                                                  190191-89-0P
    190191-90-3P
                    190191-91-4P
    RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
    study); PREP (Preparation); USES (Uses)
        (thermosensitive biodegradable polymers based on poly(ether-ester)
       block copolymers)
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was added PEG which was predried under high vacuum at an elevated temperature

50-56-6, Oxytocin, biological studies 50-76-0, Actinomycin d 5-Fluorouracil 58-82-2, Bradykinin 59-05-2, Methotrexate 1066-17-7, Colistin 1393-25-5, Secretin 1404-00-8, Mitomycin 1405-87-4, 1405-97-6, Gramicidin 1406-11-7, Polymixin 1407-47-2, Bacitracin Angiotensin 1947-37-1, Tetragastrin 5534-95-2, Pentagastrin 8011-61-8, Tyrocidine 8049-62-5, Zinc insulin 9002-60-2, Adrenocorticotrophic hormone, biological studies 9002-62-4, Prolactin, biological studies 9002-72-6, Growth hormone 9002-76-0, Gastrin 9007-12-9, Calcitonin 9007-92-5, Glucagon, biological studies 9015-94-5, Renin, biological studies 9034-39-3, Growth hormone releasing hormone 9034-40-6, Luliberin 9061-61-4, Nerve growth factor 11000-17-2, Vasopressin 11056-06-7, Bleomycin 15663-27-1, Cisplatin 20830-81-3, Daunorubicin 23214-92-8, Doxorubicin 24305-27-9, TSH releasing hormone 25316-40-9, Adriamycin 33069-62-4, Taxol 41575-94-4, Carboplatin 51110-01-1, Somatostatin 60118-07-2, Endorphin 62229-50-9, Urogastrone 81627-83-0, Macrophage colony stimulating factor 83869-56-1, Granulocyte macrophage colony stimulating factor 114977-28-5, Taxotere 143011-72-7, Granulocyte colony stimulating factor RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (thermosensitive biodegradable polymers based on poly(ether-ester) block copolymers)

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(FILE 'HOME' ENTERED AT 10:08:29 ON 11 MAR 2005)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, CANCERLIT, JAPIO' ENTERED AT 10:08:52 ON 11 MAR 2005 3460 S (ANTIGEN BINDING SITE) L1 0 S (COMPLEMENTARY DETERMING SEG?) L22 S (COMPLEMENTARY DETERMINING SEG?) L3 L40 S L1 AND L2 L5 2376516 S ANTIBOD? L6 93 S L5 AND (180 DEGREE) L7 13825 S L5 AND (HYDROPHOBIC?) L8 3 S L6 AND L7 3 DUPLICATE REMOVE L8 (0 DUPLICATES REMOVED) Ь9

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d his

(FILE 'HOME' ENTERED AT 10:08:29 ON 11 MAR 2005)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, CANCERLIT, JAPIO' ENTERED AT 10:08:52 ON 11 MAR 2005 3460 S (ANTIGEN BINDING SITE) L1 0 S (COMPLEMENTARY DETERMING SEG?) L22 S (COMPLEMENTARY DETERMINING SEG?) L3 0 S L1 AND L2 L42376516 S ANTIBOD? L5 93 S L5 AND (180 DEGREE) L6 L7 13825 S L5 AND (HYDROPHOBIC?) 3 S L6 AND L7 L8 L9 3 DUPLICATE REMOVE L8 (0 DUPLICATES REMOVED)

=>